FIG.1a

5' <u>ICT AGA CCA 166</u> TAC GTC CTG TAC AAA CCC CA 3' s' AAG CII ICA ITG ITI GCC ICC CIG CIG CGG 3 E.c. A-Gluc back:

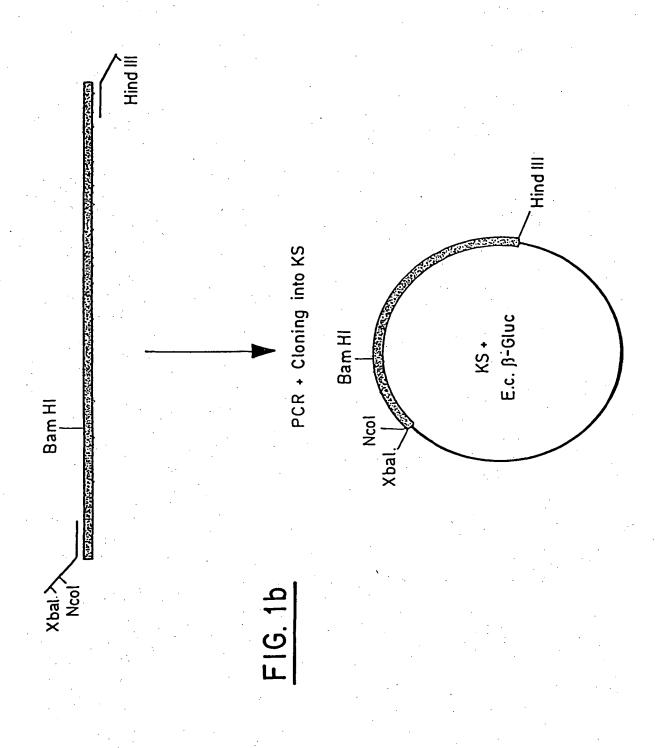


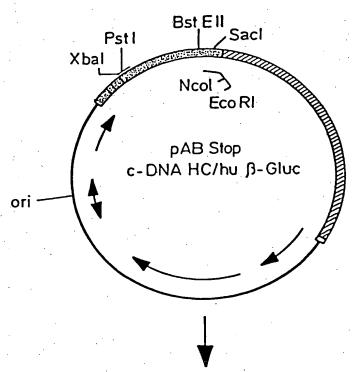
FIG. 2a

Fab HC for: 5' GAA TIC CAT GGA ACC AGA ACC AGA ACC GAG CTC AAC TCT N2 CTT GTC CAC CTT GGT GTT 3' C H 1

Fab HC back: 5' ICT AGA TAA CGA GGG CAA AAA ATG GAG GTC CAA CTG CAG N2b

GAG AGC 3'

FIG. 2b



PCR + Cloning into KS+

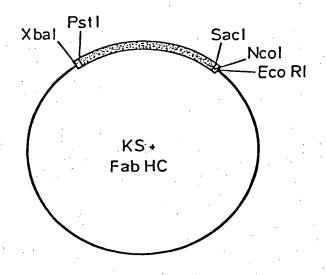
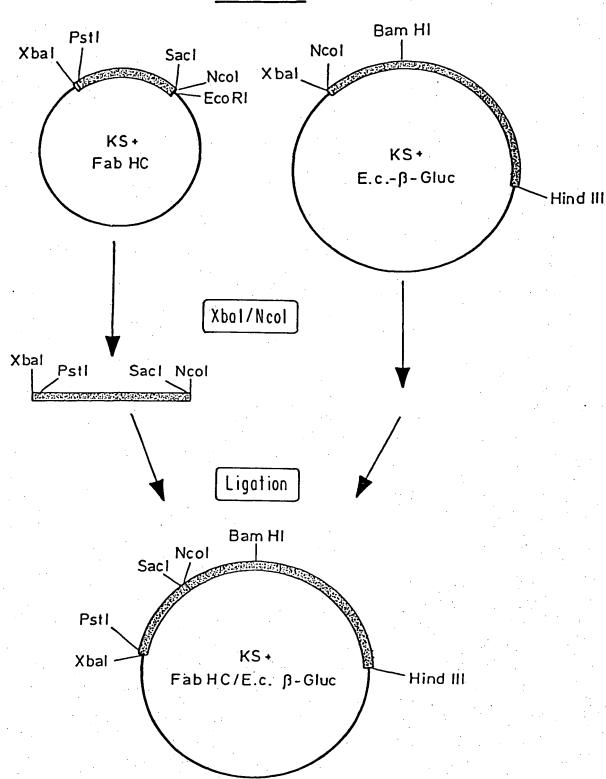
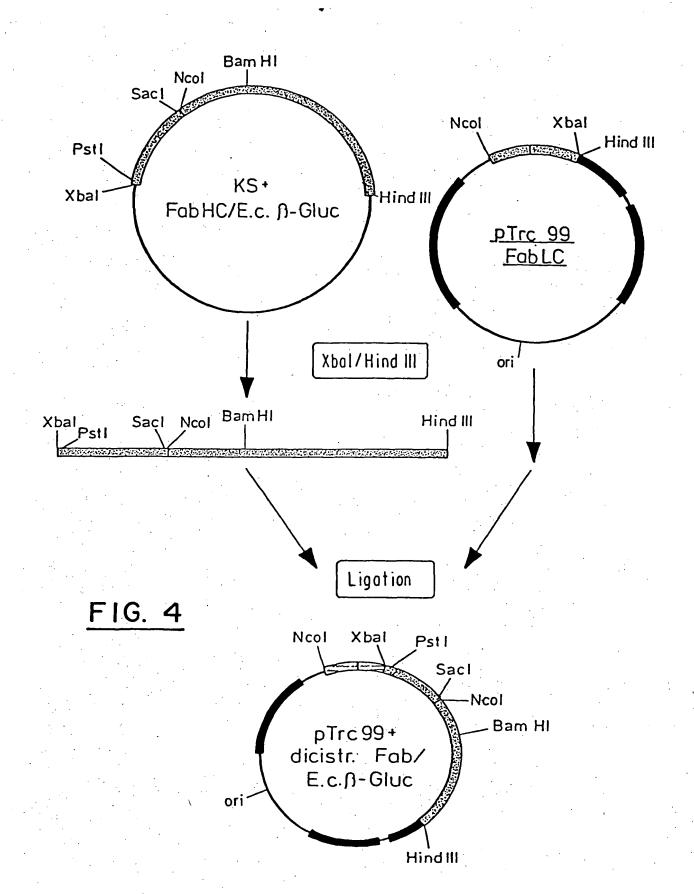
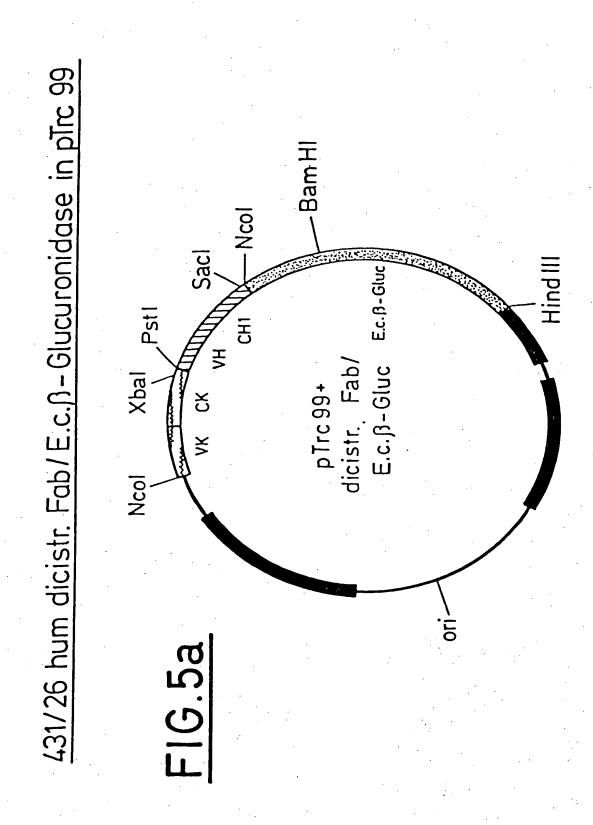


FIG.3







Attorney Docket No.: 38005-0183

5b

N S		X		
AGA A rg	AAG Lys	చ్చ క	ට වූ	CAA GI n
GAC A Asp A	CAG Gl n	GTG Va I	CTC	06 Gl √
GGT G	CAG Gl n	GGT Gly	AGC Ser	F &
GTG G Val G	TAC	TCT Ser	AGC	ACG
AGC G Ser V	TGG Trp	GCT Ala	ATC 11e	ည ၁
GCC A	CAC His	CTG	ACC	TAT
AGC G Ser A	ATG Me 1	AAC Asn	TTC	AGT Ser
CTG A Leu S	TAC	JCC Ser	ACC	AGT Ser
SS C	AGT Ser	ACA	TTC Phe	TGG Trp
AGC AGC Ser Ser	GTA Va I	AGC Ser	GAC Asp	CAG GIn
S or A	AGT Ser	TAC	ACC	CAT
AGC C	32 8	ATC 110	GGT Gly	TGC Cys
CAG A	AGC Ser	CTG	AGC GGT Sor Gly	TAC
ACC O	ACC Thr	CTG	GGT Gly	TAC
ATG A	AGT Ser	AAG Lys	AGC Ser	ACC
AG A	TGT Cys	CCA Pro	GGT Gly	GCC Ala
21 19 0	ACC	GCT Ala	& Ser	ATC 110
sp 1	ATC 11e	AAG Lys	TTC	GAC Asp
1 CC ATG GAC ATC CAG 1 Met Asp 11e Gin	60 GTG ACC ATC ACC TGT 20 Val Thr 110 Thr Cys	120 CCA GGT AAG GCT CCA 40 Pro Gly Lys Ala Pro	180 AGC AGA TTC AGC GGT 60 Ser Arg Phe Ser Gly	240 CCA GAG GAC ATC GCC 80 Pro Giu Asp IIe Ala
CC AT 1¥Me	GTG Va I	CCA Pro	AGC 88 -	S P S
-	60 20	120 40 >	180 60\$	240 80 🕨

इंट्र

GCT GCA O

GTG Val

GAA ATC AAA (

ACC ANG GTG Thr Lys Val

FIG. 50

	\aleph			
TAT Tyr	C AG	ACG	66C Gl y	AAA
TTC Phe	7CC 88.	CTC	CAG Gl	2555
AAC	AAC	ACC Thr	CAT	ACGA
AAT AAC Asn Asn	GGT. AAC. 7 Gly Asn S	Ser Sec	ACC	GATA
CTC	, g	AGC Ser	GTC	Xbal TCTAGATAACGAGGCCAAA
CTG	\$ 10 2 10	CIC	GAA Glu	TAG
TGC Cys	CIC Bu	AGC Ser	TGC	TGT
GTG Va I	GAT AAC GCC (Asp Asn Ala L	TAC	SCC Ala	GAG Gl u
GTT Va I	AAC Asn	ACC	TAC	GG. GI Y
GGA ACT GCC TCT Gly Thr Ala Ser	GAT Asp) Ser	AAA CAC AAA GTC TAC GCC Lys His Lys Val Tyr Ala	AGG GGA Arg Gly
Ala Ala	s TGG AAG GTG	Ser Lys Asp	AAA Lys	AAC Asn
ACT	AAG Lys	AAG Lys	CAC Hi s	TTC
GGA GI Y	TGG Trp	AGC Ser	AAA Lys	AGC Ser
Ser	CAG	CAG GAC GIn Asp	TAC GAG Tyr Glu	AAG Lys
AAA Lys	GTA Va i	CAG GI n	TAC	ACA Thr
TTG	AAA Lys	SAG Gi u	GAC	GTC Val
CAG Gl n	GCC Ala	ACA Thr	GCA A I a	CCC Pro
S S S C C	GAG GI U	GTC Val	AAA Lys	AGC TCG CCC GTC Ser Ser Pro Val
360 TCT GAT GAG 120 ►Ser Asp Glu	AGA A rg	AGT	CTG AGC AAA GCA GAC Leu Ser Lys Ala Asp	AGC
हिंद	00 c 0 c	GAG Glu	CTG	CTG
360 120▶	420 CCC AGA GAG GCC AAA 140 Pro Arg Glu Ala Lys	480 GAG AGT GTC ACA 160 GIU Ser Val Thr	540 (600 CTG 200▶ Leu
				•

GGT CCA GGT CTT GTG AGA CCT AGC CAG ACC CTG Gly Pro Gly Leu Val Arg Pro Ser Gln Thr Leu GAG AGC Glu Ser Pstl CTC Leu § 5 664 AA ATG GAG
1 Mot GIU

>

GTG Va I	TA THE	CAG G(n	AGA A rg	STC Val	Arg	9 s
755 9. T		AAC	S A B	ACA GTC Thr Val	325	§ 2
S E	द्धाः व्य	AAG Lys	IGT. Cys	GTC Val	TGC Cys	25 c
7 T T T	AGT.	AGC Se	TAT	Car	8 g	TTC FF.
AGC Se r	TAC	ACC THE	TAT Tyr	, See .	SCG Ala	TAC
The Cys The Val See Gly Phe The 11e See See Gly Tyr See Trp HIs Trp Val	CGA GGT CTT GAG TGG ATT GGA TAC ATA CAG TAC AGT GGT ATC Arg Gly Leu Glu Trp II.e Gly Tyr IIe Gln Tyr Ser Gly IIe	AAC CCC TCT CTC AAA AGT AGA GTG ACA ATG CTG GTA GAC ACC AGC AAG AAC Asn Pro Ser Leu Lys Ser Arg Val Thr Met Leu Val Asp Thr Ser Lys Asn	CTG AGA CTC AGC AGC GTG ACA GCC GCC GAC ACC GCG GTC TAT TAT TGT. GCA AGA Leu Arg Leu Ser Ser Val Thr Ala Ala Asp Thr Ala Val Tyr Tyr Cys Ala Arg	TAT GAT TAC CAC TGG TAC TTC GAT GTC TGG GGT CAA GGC AGC CTC Tyr Asp Tyr His Trp Tyr Phe Asp Val Trp Gly Gin Gly Ser Leu	ACC ANG GCC CCA TCG GTC ITC CCC CTG GCG CCC TGC Thr Lys Gly Pro Ser Val Phe Pro Leu Ala Pro Cys	TOT GGG GGC ACA GCG GCC CTG GGC TGC CTC AAG GAC TAC TTC CCC GAA CCG Ser Gly Gly Thr Ala Ala Leu Gly Cys Leu Val Lys Asp Tyr Phe Pro Glu Pro
g 5	ATA 116.	GTA Val	GCG Ala	₹ 5 5 5	သ ဇ	AAG
Se T	TAC	CTG	ACC	GGT GI Y	F F	GTC Val
2 % 8	ජ ර ප්	ATG Me 1	GAC Asp	7 5	CTC Val	CTC Leu
AIC 110	ATT 11.8	ACA The	A La	GTC	£ 38	TGC Cys
A F	100 Trp	STC Val	CCC Ala	GAT Asp	8 8	& © ₹
P. F.	og c og c	AGA A rg	ACT	TTC Ph.	ည္တ ဗ	CTG
3 8	CTT	AGT	GTG Va I	TAC	AAG Lys	GCC Ala
Z ž	GGT S	ANA	AGC Ser	TCG Trp	S E	A l a
\ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \	CGA A rg	CTC	AGC Ser	CAC	35. Se	A H
ž ž	\$ ` 8 8	हूं पुर	CTC	TAC	TOC TCA GCT TOC Ser Ser Ala Ser	SSC Gl √
Cys	CCA CCT (Pro Pro C	သ ရ	AGA A rg	GAT	ध क्र	866 61 ×
} <u> </u>	ည် <u>ရ</u>	AAC	CTG	TAT	£ 35	Se Ter
Leu	5 5 5 7	TAC	76C Ser	GAC Asp	GTC	ACC
20 Ser Leu	83 AGA 40 ► A rg	43 AAC 60▶Asn	03 TTC 80 € Phe	e g	A F	AGC Ser
20	783 AGA CAG 40 P A rg Gin	843 AAC 60 Asn	903 80 \	963 GAA GAC 100 Gu Asp	1023 ACA 120 Thr	1083 AGC ACC 140 Ser Thr
			•	•		

F16. 56

						ш
GTC Va I	TTC Leu	AAG Lys	CCT A 79	ATT 116	AGT Sør	င် ဗ
Ala Ala	AGC Ser	GAC Asp	AC Th	ত ও ড	ය වූ ද	TAT Tyr
88 6	AGC Ser	STG Val	85°	TGT Cys	8 g	755 7 rp
F E	2 %	AAG Lys	ACC Thr	Asn	GTG Va I	STC Val
S F	SS of	8 <u>F</u>	G C	ت کې ق	GCT A! a	AAC Asn
S E	GTG Va I	AAC Asn	GTA Va I	000 A rg	ATT 110	မှ ဗို ဗ
GTC Val	A T	AGC Ser	CCT P ro	GAT	S A	A l a
ပ္ပ ခွဲ	GTG Val	သ ဥ	CGT A rg	CTC	000 P rg	TAT Tyr
Se Se	GTG Va I	AAG Lys	GTA Va I	AGT Ser	GAA AGC GI u Ser	AAT
ACC Thr	Ser Ser	S E s	ATG Me i	71. P.	GAA Gi u	CGT A rg
GCC CTG Ala Leu	AGC Se	AST	Nool TCC ATG Ser Met	S CCA TTC	TTA CAA Leu Gin	GCC GAT GCA GAT ATT CGT AAT TAT GCG Ala Asp Ala Asp 110 Arg Asn Tyr Ala
Ala Ala	CTC	STC Val	GGT GI Y	CTG TGG Leu Trp	TTA	GAT Asp
ე დ Հ	32 %	AAC Asn	8	CTS	SCG Ala	A SCA
5 3	TAC Tyr	ACC TGC Thr Cys link	GGT GI Y	ය වේ ද	AGC	GAT Asp
Asn	CTC	ACC Thr	17. §	GAC Asp	₹ © 0	600 A a
25 F	SG. SG. Y	TAC	GGT GI y	CTC	755 7 rp	35 g
, J.	हूं दु	ACC	CTC	AAA Lys	756 T rp	CAG G
GTG Val	Se.	5 0 0	Saci GAG CTC GIU Leu	: AAA AAA Lys Lys	CGT A rg	GAT
ج آج	CAG G n	Agc Thr	GTT Va I	ATC 118	r cag cgr 166 o Gin Arg Trp	AAC
STG .	CTA Leu	ე ლ	AGA A rg	5 5 0 5	GAT	E d
1143 GTG / 160 Val	1203 CTA CAG 180⊁ Leu Gin	1263 GGC 200▶ GI y	1323 AGA 220 ^b A rg	1383 GAA ATC 240 GI u 11 e	1443 GAT 260 Asp	1503 TTT AAC GAT CAG 280 Phe Asn Asp Gin

FIG. 5f

GCG Ala	ပ္ပ 🌫 ဗ	ATC 110	S F	ATC 118	GTC	द्ध ब ≺
CAT Asp	သွ ဗ	CGT A rg	ATT 110	BamHI GGC ATC GI y 11 e		AAT Asn
TTC GAT GCG Phe Asp Ala	CAG GGC Gin Giy	GTA CGT ATC Val Arg IIe	GIG ATT ACC	SCC Ala	ACC Thr	SCC Ala
AAA GGT TGG GCA GGC CAG CGT ATC GTG CTG CGT Lys Gly Trp Ala Gly Gln Arg IIe Val Leu Arg	TOG GTC AAT AAT CAG GAA GTG ATG GAG CAT Trp Val Asn Asn Gin Giu Val Met Giu His	ACG CCG TAT GTT ATT GCC GGG AAA AGT Thr Pro Tyr Val 110 Ala Gly Lys Sor	AAC AAC GAN CTG AAC TGG CAG ACT ATC CCG CGG GGA ATG Asn Asn Giu Leu Asn Trp Gin Thr 11e Pro Pro Giy Mot	AAC GGC AAG AAA AAG CAG TCT TAC TTC CAT AAT TTC TTT AAC TAT GCC Asn Gly Lys Lys Lys Gln Ser Tyr Phe His Asn Phe Phe Asn Tyr Ala	GTA ATG CTC TAC ACG CCG AAC ACC TGG GTG GAC GAT ATC ACC GTG Val Met Leu Tyr Thr Pro Asn Thr Trp Val Asp Asp 11e Thr Val	GTC GCG CAA GAC TGT AAC CAC CCG TCT GTT GAC TGG CAG GTG GTG GCC AAT GGT Val Ala Gin Asp Cys Asn His Ala Ser Val Asp Trp Gin Val Val Ala Asn Gly
Cec City	250 0 u	AAA Lys	S S ≥ 2	ASO	GAT Asp	GTG Va I
GTG Va I	ATG Me I	ර ල ද	CCG P ro	TTT.	GAC Asp	CAG G
ATC ==	GTG Va I	A B a	CCG P ro	TTC Phe	GTG Val	700 7 rp
CGT A rg	0 P	ATT 110	ATC 119	AAT Asn	755 7.7	GAC Asp
5 5 5 5 7 8	5 g	GTT Va I	ACT	CAT His	ACC	GTT Va I
S S S	AAT Asn	TAT Tyr	CAG G n	77.C	AAC	<u>8</u>
SCA A I a	AAT	CCG Pro	755 7 rp	TAC	SSS Pro	CCG Ala
755 T.p	GTC Val	ACG	AAC	ह्य	ACG	S H
हुत् र	755 7 rp	GTC Val	CTG	CAG G n	ACC	ASD
AAA Lys	GTG Va I	GAT Asp	GAA Gi u	AAG Lys	TAC	TGT
SSS Pro	MA Lys	GAA GCC GAT GTC	AAC	AAA Lys	Cit	GAC Asp
ATA 110	25 S ⊆ Z ✓			AAG Lys	ATC Me t	3 5
TTT ATA CCG /	CAT TAC GGC AAA GTG HIS Tyr GIY Lys Val	F &	GTG Val	2 G	GTA Va I	CCC A 1 a
GTC Val	CAT	ე <u>გ</u>	TGT	AAC	AGC Ser	
2 5 5 5	ACT	ACG Thr	GTT Va I	र छ ह	CGC A rg	CAT H s
CGC A rg	GTC Val	TAT Tyr	ACC	GAC	CAT	ACG
1563 CGC GAA 300 A rg Gl u	1623 GTC 7 320 Val	1683 TAT / 340 Tyr	1743 ACC (360 Thr \	1803 GAC GAA 380⊁Asp GIu	1863 CAT (400) HIS A	1923 ACG (

FIG. 5g

ACT	TAT	ე / დ ფ	FF. Page	CTC	CAT
ပ္ပ တီ ဗီ	CTC	GTC	TAC	GTG Va I	75G
g 6	TAT	OSC A rg	TTC Phe	AAC	ACC
© & © ₹	GGT . G	CTT C	88 e e	SAT A	SGT
P.C.	G L S		AAA -ys	A TTC GAT AAC	rac o
GCA ACT Ala Thr	द्धाः अ	TGT GAT ATC TAC CCG Cys Asp IIe Tyr Pro	CAG ITC CTG AIT AAC CAC AAA Gin Phe Leu ile Asn His Lys	89 / 00 ×	GAC CAC GCA TTA ATG GAC TGG ATT GGG GCC AAC TCC TAC CGT Asp His Ala Leu Met Asp Trp 11e Gly Ala Asn Ser Tyr Arg
STT (CCG Pro	ATC .	AAC (AAA GGA Lys Gly	AAC :
CAA CAG GTG GTT GIn Gin Val Val	ខ្មុំខ្	GAT	ATT A	6 y 1	300 y
CAG G n	10G	TGT (Cys /	CTG .	CGT (6G6 61 Y 1
S C C	CTC	GAG Glu	FI G	TTA CGT Leu Arg	ATT 110
GAT Asp	E SC Eis	A THI	CAG Gln	GCG GAC	Trp
ccr car ccc car Arg Asp Ala Asp	ccc P ro	AAA AGC CAG ACA Lys Ser Gin Thr	\$ 0 0	GCG Ala/	GAC Asp
GAT Asp A	AAT	AGC	g ∨	GAT Asp	ATG Me t
CGT A	GTC Val	AAA Lys	AAG Lys	GAA GAT Glu Asp	TTA Leu
CTC	GTG Val	GCC Ala	GTG Va I	CAT His	A la
ت و ت و	C S	A F	GCA GTG AAG GGC Ala Val Lys Gly	CGT A rg	CAC
GTT Va I	TTG	GIC ACA Val Thr	GTG Val	GGT Gly	GAC
AGC Ser	ACT	TGC			CAC H s
GTC	866 Gl √	CTG	000 A rg	ე დ გ	GTC Val
GAT	AGC Ser	G CAA	ATC 110	ACT THE	ATG Me t
1983 GAT GTC 440 Asp Val	2043 AGC 460 Ser	2103 GAA CTG 480∮GIU Leu	2163 ATC CCG TCA 500 IIe Arg Ser	2223 ACT GGC TTT 520 Thr GLY Phe	2283 ATG GTG CAC 540 Met Val His
•					.*

F1G. 5F

GAT	AAG Lys	G G G	ATT 110	G CAA	GAC	र्ड ठे
ATT 119	GGC AAC AAG Gly Asn Lys	TTA	AGT Ser	GCG Ala	TGC	TAT TAC Tyr Tyr
GTG Val	ე დ დ	CAC His	TGG Trp	CTG	2 4	TAT
GTG Val	GCG Ala	GCG Ala	ATG Me I	CCA o	ATG Mo 1	ccr A rg
ATC 116	CAN GCG	CAA GCG GIn Ala	GTG ATG Val Met	GCG A l a	GTA Val	AAC
CTC GAC TGG GCA GAT GAA CAT GGC ATC GTG Leu Asp Trp Ala Asp Glu His Gly Ile Val	TTC PH e	CAG Gr	CCA AGC GTG (Pro Ser Val V	CAA GGT GCA CGG GAA TAT TTC GCG GIN GIY AIA AIG GIU TYR Phe AIA	AAT GTA ATG 1	crc AAC CGT 1 Leu Asn Arg T
CAT His	3GT Sl y		AGC Ser	TAT	STS A l	် ဒို
GAA Gi u	TTA GGC ATT (Leu Gly 119 (GCA GTC AAC GGG GAA ACT Ala Val Asn Gly Glu Thr	S S	G CAA	T CCG ATC ACC TGC C g Pro 11e Thr Cys \	TTT GAT GTG CTG Phe Asp Val Leu
GAT	SS SI →	565 GI Y	ATA GCG CGT GAC AAA AAC CAC 110 Ala Arg Asp Lys Asn His	CGG A rg	A THE	GTG Val
GCA Ala	TTA	AAC	AAC Asn	GCA Ala	ATC 119	GAT
TGG Trp	Ser	GTC	AAA Lys	GGT GI y	ეე ი ი	TTT
GAC	CEC	A SCA	GAC	S C S	CGT A rg	CTC
	GGC TTT AAC CTC Gly Phe Asn Leu	CAG Glu	CGT A rg	r ACC CGT CCG	CCG ACG CGT Pro Thr Arg	GAT
ATG Me t	Phe	G u G u	GCG Ala	CGT A rg	CCG Pro	AGC Ser
GAG Gl u	00 01 × 01 ×	AGC G	ATA 119	ACC	GAC Asp	ATC 110
n G u	GTC	TAC	nen Cuc	GAT	AAA CTC Lys Lou	GAT ACC ATC AGC GAT CTC Asp Thr 11e Ser Asp Leu
GCT Ala	GCT	CTG	GAG Gl c	CCG GAT Pro Asp	AAA Lys	GAT
TAC		G L	AAA Lys	GAA Gl u	CGT A rg	ACC
CCT Pro	ACT	AAA Lys	ATT 118	AAC	ACG	CAC His
TAC	G v	463 CCG AAA 600 Pro Lys	GCG Ala	SCC Ala	A B A	GCT Ala
2343 TAC CCT TAC 560 Tyr Pro Tyr	2403 GAA ACT GCT 580 € GIU Thr Ala	2463 CCG AAA 600 Pro Lys	2523 GCG ATT AAA 620 Ala 110 Lys	2583 GCC AAC GAA 640 A I a Asn Glu	2643 GCA ACG CGT 660 Ala Thr Arg	2703 GCT CAC ACC 680 AIs His Thr
						* •

2763 TGG TAT GTC CAA AGC GGC GAT TTG GAA ACG GCA GAG AAG GTA CTG GAA AAA GAA CTT CTG 700 Trp Tyr Val Gln Ser Gly Asp Leu, Glu Thr Ala Glu Lys Val Leu Glu Lys Glu Leu Leu

F16. 51

		_		
TTA Leu	CTC	AAT Asn	ATC 119	00 € 00 7
ACG Thr	TGG Trp	TGG Trp	555 Gl 4	ACT
GAT. Asp	A P a	GTA Va I	AAA Lys	155 Trp
GTG Va I	TGT	CAG GI n	AAG Lys	CGC A rg
. SS >	CAG Gl n	G L	AAC	AAA Lys
TAC	TAT	GGT GI Y	GGT Gly	CAA AAA CGC
GAA GI u	CAG Glu	GTC	00 √	CTG
ACC	G GA	GTC	GTT Val	CTG
ATC 1 e	AGT Ser	GCC Ala	CCC A rg	Phe
ATC 119	TGG Trp	AGC	TTG	GCT Ala
ATT ATC /	ATG Me t	GTC Val	ATA II e	GCG A I a
CCG P ro	GAC Asp	CGC A rg	03 13 13 13 13 13 13 13 13 13 13 13 13 13	70G Ser
CAG Gln	ACC Thr	GAT	S E E	AAG Lys
CAT Hi s	TAC	TTT GAT Phe Asp /	7CG 28.	CCG Pro
CIG	ATG Me i	GTC	ACC	AAA Lys
AAA Lys	S S	CGC A rg	GCG Ala	CGC A rg
GAG AAA (CAC TCA His Ser	CAC CGC His Arg	TTT Pho	GAC CGC
ପ୍ର ପ୍ର	CTC	TAT	GAT	cgc A rg
TCG Trp	SS ≥ S = S	VTG fe 1	GCC Ala	ACT
Al a	Ala Ala	GAT Asp	TTC Pre	± C
2823 GCC TGG 720▶A!a Trp	2883 GCC GGG CTG 740⊁Ala Gly Leu	2943 GAT A 760 Asp M	3003 TTC GCC GAT 780 Phe Ala Asp	3063 TTC ACT CGC 0 800 Phe Thr Arg A

Hindlil 3123 ATG AAC TTC GGT GAA AAA CCG CAG CAG GGA GGC AAA CAA TGA AGCTT 820 Met Asn Phe Gly Glu Lys Pro Gln Gln Gly Gly Lys Gln ...



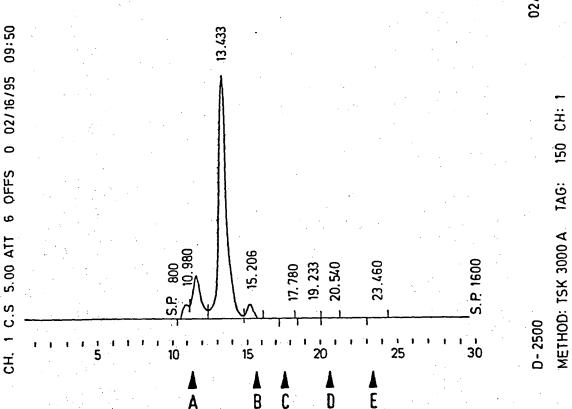


FIG.6

TSK - 3000 Gel chromatography. Retention time of the fusion protein, 13.4 min.

A: Thyroglobulin 670.000 D B: Gamma - Globulin 158.000 D

C: Ovalbumin 44.000 D

D: Myoglobulin 17.000 D

E: Vitamin B-12 1.350 D